polypeptide binds to a *patched* protein or promotes proliferation of spermatogonia, Sertoli cells, or Leydig cells.

- 39. (Reiterated) An expression vector, capable of replicating in at least one of a prokaryotic cell and eukaryotic cell, comprising the nucleic acid of claim 35.
- 40. (Reiterated) A host cell transfected with the expression vector of claim 39 and expressing said recombinant polypeptide.
- 42. (Reiterated) A recombinant transfection system, comprising
- (i) a gene construct including the nucleic acid of claim 35, operably linked to a transcriptional regulatory sequence for causing expression of the hedgehog polypeptide in eukaryotic cells, and
- (ii) a gene delivery composition for delivering said gene construct to a cell and causing the cell to be transfected with said gene construct.
- 43. (Twice Amended) The recombinant transfection system of claim 42, wherein the gene delivery composition is selected from a group consisting of a recombinant viral particle, a liposome, and a poly-cationic nucleic acid binding agent.
- 49. (Twice Amended) An isolated nucleic acid comprising a nucleotide sequence which encodes a polypeptide of SEQ ID No. 17 or an N-terminal fragment thereof having a molecular weight of about 19 kD, which polypeptide binds to a patched protein.
- 52. (Reiterated) A nucleic acid according to claim 49, further comprising a transcriptional regulatory sequence operably linked to said nucleotide sequence.
- 53. (Reiterated) An expression vector, configured for replication in at least one of a prokaryotic cell and eukaryotic cell, comprising the nucleic acid of claim 49.
- 54. (Reiterated) A host cell transfected with the expression vector of claim 53.



- 62. (Reiterated) The nucleic acid of claim 49, comprising the nucleotide sequence of SEQ ID No. 8.
- 64. (Amended) An isolated nucleic acid comprising the nucleotide sequence of SEQ ID No: 8 or its complementary nucleotide sequence.
- 66. (Amended) An expression vector comprising the nucleic acid of claim 64.
- 68. (Reiterated) The nucleic acid of claim 64, which encodes an amino acid sequence of SEQ ID NO: 17.
- 69. (Amended) A host cell transfected with the nucleic acid of claim 64 and expressing said recombinant polypeptide.
- 70. (Amended) A host cell transfected with the expression vector of claim 66 and expressing said recombinant polypeptide.
- 75. (Reiterated) The nucleic acid of claim 49, which nucleic acid encodes a polypeptide including amino acids 23-198 of SEQ ID No: 17.
- 76. (Reiterated) The nucleic acid of claim 49, which nucleic acid encodes a polypeptide including SEQ ID No: 17.
- 77. (Amended) An isolated nucleic acid encoding a polypeptide consisting essentially of a hedgehog polypeptide which is at least 98 percent identical to either SEQ ID No: 17 or an N-terminal fragment thereof having a molecular weight of about 19 kD, which hedgehog polypeptide binds to a patched protein or promotes proliferation of spermatogonia, Sertoli cells, or Leydig cells.
- 78. (Amended) An isolated nucleic acid encoding a polypeptide consisting of a hedgehog polypeptide which is at least 98 percent identical to either SEQ ID No: 17 or an N-terminal



fragment thereof having a molecular weight of about 19 kD, which hedgehog polypeptide binds to a patched protein or promotes proliferation of spermatogonia, Sertoli cells, or Leydig cells.

79. (Amended) An isolated nucleic acid encoding a polypeptide comprising a hedgehog amino acid sequence which is at least 98 percent identical to SEQ ID No: 17, which hedgehog amino acid sequence binds to a patched protein or promotes proliferation of spermatogonia, Sertoli cells, or Leydig cells.

The amended claims are re-stated below to reflect changes since the last filing.

- 35. (Thrice Amended) An isolated nucleic acid encoding a polypeptide comprising a hedgehog [amino acid sequence]polypeptide which is at least 98 percent identical to either SEQ ID No: 17 or an N-terminal fragment of SEQ ID No: 17 having a molecular weight of about 19 kD, which hedgehog [amino acid sequence]polypeptide binds to a patched protein or [regulates]promotes proliferation of [testicular germ line cells]spermatogonia, Sertoli cells, or Leydig cells.
- 49. (Twice Amended) An isolated nucleic acid comprising a nucleotide sequence which encodes [an amino acid sequence] a polypeptide of SEQ ID No. 17 or an N-terminal fragment thereof having a molecular weight of about 19 kD, which [amino acid sequence] polypeptide binds to a patched protein.
- 63. (CANCEL) An isolated nucleic acid which encodes a naturally occurring Desert hedgehog protein of human origin or an N-terminal fragment thereof, having a molecular weight of about 19 kD.
- 64. (Amended) [The nucleic acid of claim 63, which contains either] An isolated nucleic acid comprising the nucleotide sequence of SEQ ID No: 8 or its complementary nucleotide sequence.



- 65. (CANCEL) The nucleic acid of claim 63, which is inserted into an expression vector.
- 66. (Amended) [The nucleic acid of claim 64, which is inserted into an] An expression vector comprising the nucleic acid of claim 64.
- 67. (CANCEL) The nucleic acid of claim 63, which encodes an amino acid sequence of SEQ ID NO: 17.
- 69. (Amended) [The nucleic acid] A host cell transfected with the nucleic acid of claim 64 and expressing said recombinant polypeptide [63 or 64, which is introduced into an appropriate host].
- 70. (Amended) [The nucleic acid] A host cell transfected with the expression vector of claim 66 and expressing said recombinant polypeptide [65 or 66, which is introduced into an appropriate host].
- 77. (Amended) An isolated nucleic acid encoding a polypeptide consisting essentially of a hedgehog [amino acid sequence]polypeptide which is at least 98 percent identical to either SEQ ID No: 17 or an N-terminal fragment thereof having a molecular weight of about 19 kD, which hedgehog [amino acid sequence]polypeptide binds to a patched protein or [regulates]promotes proliferation of [testicular germ line cells]spermatogonia, Sertoli cells, or Leydig cells.
- 78. (Amended) An isolated nucleic acid encoding a polypeptide consisting of a hedgehog [amino acid sequence]polypeptide which is at least 98 percent identical to either SEQ ID No: 17 or an N-terminal fragment thereof having a molecular weight of about 19 kD, which hedgehog [amino acid sequence]polypeptide binds to a patched protein or [regulates]promotes proliferation of [testicular germ line cells]spermatogonia, Sertoli cells, or Leydig cells.
- 79. (Amended) An isolated nucleic acid encoding a polypeptide comprising a hedgehog amino acid sequence which is at least 98 percent identical to SEQ ID No: 17, which hedgehog amino acid sequence binds to a patched protein or [regulates] promotes proliferation of [testicular germ line cells] spermatogonia, Sertoli cells, or Leydig cells.